## **IN THE CLAIMS:**

1. (Currently amended) A method for detecting a variant HBV which exhibits an altered sensitivity to an agent, said method comprising:

generating a genetic construct comprising a replication competent amount of the genome from said variant HBV contained in or fused to an amount of a baculovirus genome capable to infect cells of hepatic origin and then infecting said cells with said construct;

contacting said cells, before, and/or during, and/or after infection, with the agent to be tested;

optionally further infecting said cells with the same genetic construct or a genetic construct comprising the genome of HBV wild type or another HBV variant;

culturing said cells for a time and under conditions sufficient for the variant HBV to be detected to replicate, express genetic sequences and/or or assemble and/or or release virus or virus-like particles if resistant to said agent; and

subjecting the cells, cell lysates or culture supernatant fluid to viral- or viral-component-detection means to determine whether or not the variant <u>HBV</u> virus has replicated, expressed genetic material, and/or assembled, and/or or been released in the presence of said agent.

- 2. (Original) A method according to Claim 1 wherein the variant HBV is capable of replicating in the presence of an agent which inhibits or reduces infection, replication or assembly of a reference HBV.
- 3. (Original) A method according to Claim 2 wherein the agent is a nucleoside analogue or a non-nucleoside analogue.
- 4. (Currently amended) A method according to Claim 3 wherein the agent is a non-nucleoside analogue reverse transcriptase inhibitor, and/or a non-nucleoside analogue DNA dependent DNA polymerase inhibitor, or both a non-nucleoside analogue reverse transcriptase inhibitor and a non-nucleoside analogue DNA dependent DNA polymerase inhibitor.

- 5. (Original) A method according to Claim 3 wherein the nucleoside analogue is 3TC, PMEA or PCV.
- 6. (Original) A method according to Claim 2 wherein the agent is an immunointeractive molecule.
- 7. (Original) A method according to Claim 6 wherein the immunointeractive molecule is an antibody.
- 8. (Currently amended) A method according to any one of Claims 1 to 7 wherein the variant HBV comprises an altered HBV DNA polymerase, an altered HBV precore promoter or basal core promoter, an altered HBsAg, or a combination thereof.
- 9. (Currently amended) A method according to Claim 7 8 wherein the altered HBB DNA polymerase is selected from the group consisting of L426I/V, L428I/VN480G, N485K, K495R, R499Q, G499E, W499Q, F512L, I515L, V519L, L526M, M550V, M550I, V5531, S565P.
- 10. (Currently amended) A method according to any one of Claims 1 to 7 8 wherein the altered variant HBV is a multiple mutant selected from the group consisting of L526M/M5501, L526M/M550V, V519L/L526M/M550V and V519L/L526M/M5501.

## 11. (Canceled)

- 12. (Currently amended) A method according to Claim 11 8 wherein the altered HBV precore promoter or basal core promoter is selected from the group consisting of A1814T, C1856T, G1896A, G1897A, G1898A, G1899A, G1896A/ G1899A, A1762T/G1764A, T1753C, G1757A and C1653T (where the numbering is from the unique *Eco*R1 site in HBV).
  - 13. (Canceled)

- 14. (Currently amended) A method according to Claim 13 8 wherein the altered HBsAg is selected from the group consisting of G112R, T123P Y/F134S, D144E, G145R, A157D, E164D, F170L, M195I, W196L, W196S, W196STOP, M198I, W199S, S204T, S210R.
- 15. (Currently amended) A method according to Claim 14 8 wherein the altered HBsAg is selected from the group consisting of D144E, G145R, A157D, E164D, M195I, W196L, W196L, W196S, W196STOP, M198I, W199S and S210R.

## 16-19. (Canceled)

- 20. (Currently amended) An HBV variant or a recombinant or derivative form thereof or a chemical equivalent thereof or a recombinant or chemical equivalent of a component thereof detected by the method according to any one of Claims 1 to 19 Claim 8.
- 21. (Currently amended) A method according to any one of Claims 1 to 7 or any one of Claims 8 to 19 Claim 8 wherein the cells are co-infected with multiple combinations of the variant HBV comprises an altered HBV precore promoter or basal core promoter and/or or an altered HBV DNA polymerase or combinations thereof.
- 22. (Currently amended) A method according to any one of Claims 1 to 7 or any one of Claims 8 to 19 Claim 8 wherein the cells are superinfected with multiple combinations of the variant HBV comprising an altered HBV precore promoter or basal core promoter and/or or an altered HBV DNA polymerase or combinations thereof.
- 23. (Currently amended) A method for detecting a variant HBV comprising DNA polymerase which exhibits an altered sensitivity to an agent said method comprising:

generating a genetic construct comprising a replication competent amount of the genome from said variant HBV contained in or fused to an amount of a baculovirus genome capable to infect cells of hepatic origin and then infecting said cells with said construct;

contacting said cells, before, and/or during, and/or after infection, with the agent to be tested;

optionally further infecting said cells with the same genetic construct or a genetic construct comprising the genome of HBV wild type or another HBV variant;

culturing said cells for a time and under conditions sufficient for the variant HBV to be detected to replicate, express genetic sequences, and/or assemble, and/or release virus or virus-like particles if resistant to said agent; and

subjecting the cells, cell lysates or culture supernatant fluid or virus purified therefrom to HBV DNA polymerase assay in the presence or absence of nucleoside triphosphate analogues or non-nucleoside analogues analogue reverse transcriptase inhibitors or non-nucleoside analogues analogue DNA dependent DNA polymerase inhibitors.

24. (Currently amended) A method for detecting <u>HBV</u> DNA polymerase activity in the presence of an antiviral agent said method comprising:

generating a genetic construct comprising a replication competent amount of a genome from an HBV capable of producing said DNA polymerase, said genome contained in or fused to an amount of a baculovirus genome capable to infect cells of hepatic origin and then infecting said cells with said construct;

contacting said cells, before, and/or during, and/or after infection, with the agent to be tested;

optionally further infecting said cells with the same genetic construct or a genetic construct comprising the genome of HBV wild type or another HBV strain;

culturing said cells for a time and under conditions sufficient for the HBV and to replicate, express genetic sequences, and/or assemble, and/or release virus or virus-like particles if resistant to said agent; and

subjecting the cells, cell lysates or culture supernatant fluid or virus purified therefrom to <u>an</u> HBV DNA polymerase assay in the presence or absence of nucleoside triphosphate analogues or non-nucleoside <u>analogues</u> analogue reverse transcriptase inhibitors or non-nucleoside <u>analogues</u> analogue DNA dependent DNA polymerase inhibitors.